

**WEST**

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L7: Entry 7 of 14

File: USPT

Nov 23, 1999

DOCUMENT-IDENTIFIER: US 5989904 A  
TITLE: Selective inhibition of internally initiated RNA translation

## DEPR:

In addition, the present disclosure enables modifications of host cells to inhibit expression or activity of I-RNA. In the first instance, introduction of such modifications will determine whether I-RNA activity is essential for survival of host cells which express such I-RNAs. In one approach, an RNA having a sequence complementary to the I-RNA (i.e., an "antisense I-RNA") is expressed in a host cell (e.g., yeast) which expresses an I-RNA molecule. The vector for expression of the antisense I-RNA contains a selectable marker gene (e.g., URA 3) to ensure that only transformed cells are recovered. If no transformed cells expressing antisense I-RNA are recovered, inducible expression constructs may be tested to determine whether the antisense RNA vector can be transformed into the cell in absence of antisense I-RNA expression and whether subsequent expression inhibits any cellular function(s). Alternatively, the I-RNA gene may be eliminated using gene "knock out" methodology known in the art. For instance, in yeast cells exogenous DNA introduced into the cell efficiently and stably integrates into chromosomal DNA by homologous recombination, allowing efficient replacement of a wildtype gene with a non-functional copy. Typically, the non-functional copy is generated by replacing wildtype coding sequences with a selectable marker gene (e.g., LEU or URA). Transformation of diploid cells may circumvent possible lethal effects if some I-RNA activity is required for cell viability. Yeast or other I-RNA-expressing host cells, or extracts thereof, which have reduced I-RNA activity as a result of either an antisense or gene knock out modification according to the invention, are useful for expression of mRNAs requiring IRES-dependent translation initiation. Also contemplated are yeast or other I-RNA host cells which can be modified by gene knockout methodologies, as known in the art, to remove the gene encoding La or homologs thereof to produce host strains that are permissive for expression of proteins whose synthesis is dependent upon internal initiation of translation.

## DEPR:

Theoretically, the inhibitor RNA could inhibit IRES-dependent translation by two possible mechanisms: binding to UTR sequences as an antisense RNA or binding to protein factors needed for internal entry of ribosomes. To distinguish these two mechanisms, uniformly <sup>32</sup>P-labeled inhibitor RNA probe

two mechanisms, uniformly  $^{32}$ P-labeled inhibitor RNA probe was prepared and mixed with HeLa S10 extracts, and the resulting RNA-protein complexes were analyzed by nondenaturing polyacrylamide gel electrophoresis.

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Generate Collection

L7: Entry 9 of 14

File: USPT

Sep 21, 1999

DOCUMENT-IDENTIFIER: US 5955318 A  
TITLE: Reagents and methods useful for controlling the translation of hepatitis GBV proteins

## BSPR:

It therefore would be advantageous to provide reagents and methods for controlling the translation of HGBV proteins from HGBV nucleic acids. Such reagents would comprise antisense nucleic acid sequences or other compound which may specifically destabilize (or stabilize) the IRES structure. Such nucleic acid sequences or compounds could greatly enhance the ability of the medical community to provide a means for treating an individual infected with GB virus(es). In addition, IRESs are among the most highly conserved nucleotide sequences. Identification of such a sequence immediately suggests a target for probe-based detection reagents. Diagnostic or screening tests developed from these reagents could provide a safer blood and organ supply by helping to eliminate GBV in these blood and organ donations, and could provide a better understanding of the prevalence of HGBV in the population, epidemiology of the disease caused by HGBV and the prognosis of infected individuals. Additionally, these conserved structures may provide a means for purifying GBV proteins for use in diagnostic assays.

LIGHT set on as ''

**HILIGHT set on as ''**

? begin 5,6,55,154,155,155,156,312,39,biotech,biosci

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**Search Results -**

Terms	Documents
IRES near10 antisense\$	14

Database:

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 US Pre-Grant Publication Full-Text Database  
 JPO Abstracts Database  
 EPO Abstracts Database  
 Derwent World Patents Index  
 IBM Technical Disclosure Bulletins

IRES near10 antisense\$

Refine Search:

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**Search History****Today's Date: 9/6/2001**

<u>DB Name</u>	<u>Query</u>	<u>Hit Count</u>	<u>Set Name</u>
USPT,PGPB,JPAB,EPAB,DWPI,TDBD	IRES near10 antisense\$	14	<u>L7</u>
USPT,PGPB,JPAB,EPAB,DWPI,TDBD	IRES near5 antisense\$	7	<u>L6</u>
USPT,PGPB,JPAB,EPAB,DWPI,TDBD	11 and inhibit\$ near5 IRES near10 reporter\$	1	<u>L5</u>
USPT,PGPB,JPAB,EPAB,DWPI,TDBD	11 and inhibit\$ near5 IRES	1	<u>L4</u>
USPT,PGPB,JPAB,EPAB,DWPI,TDBD	11 and antisense near5 IRES	0	<u>L3</u>
USPT,PGPB,JPAB,EPAB,DWPI,TDBD	11 and antisense	1	<u>L2</u>
USPT,PGPB,JPAB,EPAB,DWPI,TDBD	5989904 [pn]	2	<u>L1</u>

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starting with: INHIBIT\$(INHIBIT-INPUT).P28-P87,P89-P89,P23-P27,P20-P22,P1-P18,P19-P19.

**Search Results -**

Terms	Documents
IRES near10 inhibit\$ and antisense\$	13

Database:

US Patents Full-Text Database  
 US Pre-Grant Publication Full-Text Database  
 JPO Abstracts Database  
 EPO Abstracts Database  
 Derwent World Patents Index  
 IBM Technical Disclosure Bulletins

IRES near10 inhibit\$ and antisense\$

Refine Search:

Clear

**Search History**

Today's Date: 9/6/2001

<u>DB Name</u>	<u>Query</u>	<u>Hit Count</u>	<u>Set Name</u>
USPT,PGPB,JPAB,EPAB,DWPI,TDBD	IRES near10 inhibit\$ and antisense\$	13	<u>L8</u>
USPT,PGPB,JPAB,EPAB,DWPI,TDBD	IRES near10 antisense\$	14	<u>L7</u>
USPT,PGPB,JPAB,EPAB,DWPI,TDBD	IRES near5 antisense\$	7	<u>L6</u>
USPT,PGPB,JPAB,EPAB,DWPI,TDBD	11 and inhibit\$ near5 IRES near10 reporter\$	1	<u>L5</u>
USPT,PGPB,JPAB,EPAB,DWPI,TDBD	11 and inhibit\$ near5 IRES	1	<u>L4</u>
USPT,PGPB,JPAB,EPAB,DWPI,TDBD	11 and antisense near5 IRES	0	<u>L3</u>
USPT,PGPB,JPAB,EPAB,DWPI,TDBD	11 and antisense	1	<u>L2</u>
USPT,PGPB,JPAB,EPAB,DWPI,TDBD	5989904 [pn]	2	<u>L1</u>



Set Items Description

? s hepatitis (n) A or HAV  
 Processing  
 Processing  
 Processing  
 Processed 10 of 38 files ...  
 Processing  
 Processing  
 Processed 20 of 38 files ...  
 Processing  
 Processed 30 of 38 files ...  
 Completed processing all files

610468 HEPATITIS  
 61789038 A  
 58574 HEPATITIS(N)A  
 12689 HAV

s1 60916 HEPATITIS (N) A OR HAV

? s s1 and antisense

60916 S1  
 141760 ANTISENSE

s2 109 S1 AND ANTISENSE

? s s2 and IRES

109 S2  
 5871 IRES

s3 3 S2 AND IRES

? d s3/3/1-3

Display 3/3/1 (Item 1 from file: 34)

DIALOG(R)File 34:SciSearch(R) Cited Ref Sci

(c) 2001 Inst for Sci Info. All rts. reserv.

05540868 Genuine Article#: WF234 No. References: 45

Title: In vitro mutational and inhibitory analysis of the cis-acting  
 translational elements within the 5' untranslated region of  
 coxsackievirus B3: Potential targets for antiviral action of  
**antisense** oligomers

Author(s): Yang DC (REPRINT) ; Wilson JE; Anderson DR; Bohunek L; Cordeiro  
 C; Kandolf R; McManus BM

Corporate Source: UNIV BRITISH COLUMBIA, ST PAULS HOSP, DEPT PATHOL & LAB  
 MED, CARDIOVASC RES LAB, MCDONALD RES WING/VANCOUVER/BC V6Z 1Y6/CANADA/  
 (REPRINT); UNIV TUBINGEN, INST PATHOL/D-72074 TUBINGEN//GERMANY/

Journal: VIROLOGY, 1997, V228, N1 (FEB 3), P63-73

ISSN: 0042-6822 Publication date: 19970203

Publisher: ACADEMIC PRESS INC JNL-COMP SUBSCRIPTIONS, 525 B ST, STE 1900,  
 SAN DIEGO, CA 92101-4495

Language: English Document Type: ARTICLE (ABSTRACT AVAILABLE)

- end of record -

?

Display 3/3/2 (Item 1 from file: 98)

DIALOG(F)File 98:General Sci Abs/Full-Text

(c) 2001 The HW Wilson Co. All rts. reserv.

24045917 H.W. WILSON RECORD NUMBER: BGS199045917 (USE FORMAT 7 FOR  
 FULLTEXT)

eIF4 initiation factors: effectors of mRNA recruitment to ribosomes and  
 regulators of translation.

Gingras, Anne-Claude

Raught, Brian; Sonenberg, Nahum



Annual Review of Biochemistry v. 68 (1999) p. 913-63  
SPECIAL FEATURES: bibl il ISSN: 0066-4154  
LANGUAGE: English  
COUNTRY OF PUBLICATION: United States  
WORD COUNT: 21787

- end of record -

?

Display 3/3/3 (Item 1 from file: 399)  
DIALOG(R)File 399:CA SEARCH(R)  
(c) 2001 AMERICAN CHEMICAL SOCIETY. All rts. reserv.

122123089 CA: 122(11)123089f PATENT  
Methods for screening compounds for potential inhibitors of translation  
of viral RNAs for use as antiviral agents  
INVENTOR(AUTHOR): Miles, Vincent J.; Mathews, Michael B.; Katze, Michael  
G.; Witherell, Gary; Watson, Julia C.  
LOCATION: USA  
ASSIGNEE: Ribogene, Inc.  
PATENT: PCT International ; WO 9423041 A2 DATE: 941013  
APPLICATION: WO 94US3623 (940401) \*US 42024 (930402)  
PAGES: 194 pp. CODEN: PIXXD2 LANGUAGE: English CLASS: C12N-015/64A;  
C12N-015/62B; C12N-015/81B; C12N-015/85B; C12N-015/11B; C12N-009/12B;  
C12N-001/19B; C12Q-001/48B; C12Q-001/68B; A61K-048/00B  
DESIGNATED COUNTRIES: AT; AU; BB; BG; BR; BY; CA; CH; CN; CZ; DE; DK; ES;  
FI; GB; HU; JP; KP; KR; KZ; LK; LU; LV; MG; MN; MW; NL; NO; NZ; PL; PT; RO;  
RU; SD; SE; SK; UA; UZ; VN DESIGNATED REGIONAL: AT; BE; CH; DE; DK; ES; FR  
; GB; GR; IE; IT; LU; MC; NL; PT; SE; BF; BJ; CF; CG; CI; CM; GA; GN; ML;

-more-

Display 3/3/3 (Item 1 from file: 399)  
DIALOG(R)File 399:CA SEARCH(R)  
(c) 2001 AMERICAN CHEMICAL SOCIETY. All rts. reserv.  
MR; NE; SN; TD; TG

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? rd s2

...examined 50 records (50)  
>>>Record 266:269585 ignored; incomplete bibliographic data, not retained -  
in RD set

...examined 50 records (100)  
...completed examining records  
S4 58 RD S2 (unique items)

? d s4/3/1-58

Display 4/3/1 (Item 1 from file: 5)  
DIALOG(P)File 5:BIOSIS Previews(R)  
(c) 2001 BIOSIS. All rts. reserv.

13127319 BIOSIS NO.: 200100334468  
Relationships between the activities in vitro and in vivo of various kinds  
of ribozyme and their intracellular localization in mammalian cells.  
AUTHOR: Kato Yoshio; Kuwabara Tomoko; Warashina Masaki; Toda Hirofumi;  
Taira Kazunari(a)  
AUTHOR ADDRESS: (a)Dept. of Chemistry and Biotechnology, Graduate School of  
Engineering, University of Tokyo, Hongo, Tokyo, 113-8656:  
taira@chembio.t.u-tokyo.ac.jp\*\*Japan  
JOURNAL: Journal of Biological Chemistry 276 (18):p15378-15385 May 4, 2001  
MEDIUM: print

ISSN: 0021-9258  
DOCUMENT TYPE: Article  
RECORD TYPE: Abstract  
LANGUAGE: English  
SUMMARY LANGUAGE: English

- end of record -

?

Display 4/3/2 (Item 2 from file: 5)  
DIALOG(R)File 5: Biosis Previews(R)  
(c) 2001 BIOSIS. All rts. reserv.

10562672 BIOSIS NO.: 199699183817  
Characterization of cell lines allowing tightly regulated expression of  
hepatitis C virus core protein.  
AUTHOR: Moradpour Darius; Englert Christoph; Wakita Takaji; Wands Jack R(a)  
AUTHOR ADDRESS: (a)Molecular Hepatol. Lab., MGH Cancer Cent., 149 13th St.,  
Charlestown, MA 02129\*\*USA  
JOURNAL: Virology 222 (1):p51-63 1996  
ISSN: 0042-6822  
DOCUMENT TYPE: Article  
RECORD TYPE: Abstract  
LANGUAGE: English

- end of record -

?

Display 4/3/3 (Item 3 from file: 5)  
DIALOG(R)File 5: Biosis Previews(R)  
(c) 2001 BIOSIS. All rts. reserv.

10383901 BIOSIS NO.: 199699005046  
Amplification of the full-length **hepatitis A** virus genome by  
long reverse transcription-PCR and transcription of infectious RNA  
directly from the amplicon.  
AUTHOR: Tellier Raymond; Bukh Jens; Emerson Suzanne U; Purcell Robert H(a)  
AUTHOR ADDRESS: (a)Hepatitis Viruses Section, Lab. Infectious Diseases,  
Natl. Inst. Allergy Infectious Diseases, Natl\*\*USA  
JOURNAL: Proceedings of the National Academy of Sciences of the United  
States of America 93 (9):p4370-4373 1996  
ISSN: 0027-8424  
DOCUMENT TYPE: Article  
RECORD TYPE: Abstract  
LANGUAGE: English

- end of record -

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Display 4/3/4 (Item 4 from file: 5)  
DIALOG(R)File 5: Biosis Previews(R)  
(c) 2001 BIOSIS. All rts. reserv.

10221038 BIOSIS NO.: 199698675956  
In vivo inhibition of hepatitis B virus gene expression by **antisense**  
phosphorothioate oligonucleotides.  
AUTHOR: Moriya Kyoji; Matsukura Makoto; Kurokawa Kiyoshi; Koike Kazuhiko(a)  
AUTHOR ADDRESS: (a)First Dep. Internal Med., Fac. Med., Univ. Tokyo, 7-3-1  
Hongo, Bunkyo-ku, Tokyo 113\*\*Japan  
JOURNAL: Biochemical and Biophysical Research Communications 218 (1):p  
217-223 1996  
ISSN: 0006-291X

DOCUMENT TYPE: Article  
RECORD TYPE: Abstract  
LANGUAGE: English

- end of record -

?

Display 4/3/5 (Item 5 from file: 5)  
DIALOG(R)File 5:Biosis Previews(R)  
(c) 2001 BIOSIS. All rts. reserv.

09321307 BIOSIS NO.: 199497329677  
In situ hybridization studies of **hepatitis A** viral RNA in  
patients with acute **hepatitis A**.  
AUTHOR: Taylor Michael(a); Goldin Robert D; Ladva Suresh; Scheuer Peter J;  
Thomas Howard C  
AUTHOR ADDRESS: (a)Dep. Histopathol., St. Mary's Hosp. Med. Sch., Norfolk  
Place, Paddington, London W2 1PG\*\*UK  
JOURNAL: Journal of Hepatology 20 (3):p380-387 1994  
ISSN: 0168-8278  
DOCUMENT TYPE: Article  
RECORD TYPE: Abstract  
LANGUAGE: English

- end of record -

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Display 4/3/6 (Item 6 from file: 5)  
DIALOG(R)File 5:Biosis Previews(R)  
(c) 2001 BIOSIS. All rts. reserv.

08936189 BIOSIS NO.: 199396087690  
One-step RNA polymerase chain reaction for detection of hepatitis C virus  
RNA.  
AUTHOR: Hu Ke-Qin; Yu Chang-Hong; Vierling John M(a)  
AUTHOR ADDRESS: (a)Cedars-Sinai Med. Center, Hepatol., Suite 7511, 8700  
Beverly Blvd., Los Angeles, CA 90048\*\*USA  
JOURNAL: Hepatology 18 (2):p270-274 1993  
ISSN: 0270-9139  
DOCUMENT TYPE: Article  
RECORD TYPE: Abstract  
LANGUAGE: English

- end of record -

?

Display 4/3/7 (Item 7 from file: 5)  
DIALOG(R)File 5:Biosis Previews(R)  
(c) 2001 BIOSIS. All rts. reserv.

08404275 BIOSIS NO.: 000094121929  
IN-SITU HYBRIDIZATION STUDIES IN **HEPATITIS A** INFECTION  
AUTHOR: TAYLOR G M; GOLDIN R D; KARAYIANNIS P; THOMAS H C  
AUTHOR ADDRESS: DEP. HISTOPATHOLOGY, ST. MARY'S HOSP. MED. SCH., NORFOLK  
PLACE, PADDINGTON, LONDON W2 1PG, UK.  
JOURNAL: HEPATOLOGY 16 (3). 1992. 642-648. 1992  
FULL JOURNAL NAME: HEPATOLOGY (Baltimore)  
CODEN: HPTLD  
RECORD TYPE: Abstract  
LANGUAGE: ENGLISH

- end of record -

?

Display 4/3/8 (Item 8 from file: 5)  
DIALOG(R)File 5: Biosis Previews(R)  
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06747477 BIOSIS NO.: 000088056908  
EXPRESSION OF **A HEPATITIS B** VIRUS TRANSCRIPT CONTAINING FUSED  
MITOCHONDRIAL-LIKE DOMAINS IN HUMAN HEPATOMA CELLS  
AUTHOR: KOCH I; HOF SCHNEIDER P H; KOSHY R  
AUTHOR ADDRESS: MAX-PLANCK-INST. FUER BIOCHEMIE, MARTINSRIED BEI MUENCHEN,  
WEST GERMANY.  
JOURNAL: VIROLOGY 170 (2). 1989. 591-594. 1989  
FULL JOURNAL NAME: Virology  
CODEN: VIRLA  
RECORD TYPE: Abstract  
LANGUAGE: ENGLISH

- end of record -

?

Display 4/3/9 (Item 1 from file: 154)  
DIALOG(R)File 154: Medline(R)

07911017 93119276 PMID: 8380325  
Proteins specifically binding to the 3' untranslated region of  
**hepatitis A** virus RNA in persistently infected cells.  
Nuesch JP; Weitz M; Siegl G  
Yale New Haven Hospital, Connecticut.  
Archives of virology (AUSTRIA) 1993, 128 (1-2) p65-79, ISSN  
0304-8608 Journal Code: 8L7  
Languages: ENGLISH  
Document type: Journal Article  
Record type: Completed

- end of record -

?

Display 4/3/10 (Item 2 from file: 154)  
DIALOG(R)File 154: Medline(R)

06999644 92185479 PMID: 1312125  
Typing hepatitis C virus by polymerase chain reaction with type-specific  
primers: application to clinical surveys and tracing infectious sources.  
Okamoto H; Sugiyama Y; Okada S; Kurai K; Akahane Y; Sugai Y; Tanaka T;  
Sato K; Tsuda F; Miyakawa Y; et al  
Immunology Division, Jichi Medical School, Tochigi-Ken, Japan.  
Journal of general virology (ENGLAND) Mar 1992, 73 ( Pt 3) p673-9,  
ISSN 0022-1317 Journal Code: I9B  
Languages: ENGLISH  
Document type: Journal Article  
Record type: Completed

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?

Display 4/3/11 (Item 1 from file: 34)  
DIALOG(R)File 34: SciSearch(R) Cited Ref Sci  
(c) 2001 Inst for Sci Info. All rts. reserv.

08118163 Genuine Article#: 248HB No. References: 69  
Title: Hepatitis C virus: an overview of current approaches and progress  
Author(s): Walker MA (REPRINT)

Corporate Source: BRISTOL MYERS SQUIBB CO, PHARMACEUT RES INST, 5 RES  
PKWY/WALLINGFORD//CT/06492 (REPRINT)  
Journal: DRUG DISCOVERY TODAY, 1999, V4, N11 (NOV), P518-529  
ISSN: 1359-6446 Publication date: 19991100  
Publisher: ELSEVIER SCI LTD, THE BOULEVARD, LANGFORD LANE, KIDLINGTON,  
OXFORD OX5 1GB, OXON, ENGLAND  
Language: English Document Type: REVIEW (ABSTRACT AVAILABLE)

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Display 4/3/12 (Item 2 from file: 34)  
DIALOG(R)File 34:SciSearch(R) Cited Ref Sci  
(c) 2001 Inst for Sci Info. All rts. reserv.

05854109 Genuine Article#: XB883 No. References: 27  
Title: Properties of hepatitis delta virus ribozyme, which consists of  
three RNA oligomer strands  
Author(s): Sakamoto T; Tanaka Y; Kuwabara T; Kim MH; Kurihara Y; Katahira M  
; Uesugi S (REPRINT)  
Corporate Source: YOKOHAMA NATL UNIV, FAC ENGN, DEPT BIOENGN, HODOGAYA KU,  
79-5 TOKIWADAI/YOKOHAMA/KANAGAWA 240/JAPAN/ (REPRINT); YOKOHAMA NATL  
UNIV, FAC ENGN, DEPT BIOENGN, HODOGAYA KU/YOKOHAMA/KANAGAWA 240/JAPAN/  
Journal: JOURNAL OF BIOCHEMISTRY, 1997, V121, N6 (JUN), P1123-1128  
ISSN: 0021-924X Publication date: 19970600  
Publisher: JAPANESE BIOCHEMICAL SOC, ISHIKAWA BLDG-3F, 25-16 HONGO-5-CHOME,  
BUNKYO-KU, TOKYO 113, JAPAN  
Language: English Document Type: ARTICLE (ABSTRACT AVAILABLE)

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Display 4/3/13 (Item 3 from file: 34)  
DIALOG(R)File 34:SciSearch(R) Cited Ref Sci  
(c) 2001 Inst for Sci Info. All rts. reserv.

05598460 Genuine Article#: WJ848 No. References: 44  
Title: A one-tube method of reverse Transcription-PCR to efficiently  
amplify a 3-kilobase region from the RNA polymerase gene to the poly(A)  
tail of small round-structured viruses (Norwalk-like viruses)  
Author(s): Ando T (REPRINT) ; Monroe SS; Noel JS; Glass RI  
Corporate Source: CTR DIS CONTROL & PREVENT, VIRAL GASTROENTERITIS SECT, DIV  
VIRAL & RICKETTSIAL DIS/ATLANTA//GA/30333 (REPRINT)  
Journal: JOURNAL OF CLINICAL MICROBIOLOGY, 1997, V35, N3 (MAR), P570-577  
ISSN: 0095-1137 Publication date: 19970300  
Publisher: AMER SOC MICROBIOLOGY, 1325 MASSACHUSETTS AVENUE, NW,  
WASHINGTON, DC 20005-4171  
Language: English Document Type: ARTICLE (ABSTRACT AVAILABLE)

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Display 4/3/14 (Item 4 from file: 34)  
DIALOG(R)File 34:SciSearch(R) Cited Ref Sci  
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05540869 Genuine Article#: WF234 No. References: 45  
Title: In vitro mutational and inhibitory analysis of the cis-acting  
translational elements within the 5' untranslated region of  
coxsackievirus B3: Potential targets for antiviral action of  
**antisense** oligomers  
Author(s): Yang DC (REPRINT) ; Wilson JE; Anderson DR; Bohunek L; Cordeiro

C; Kandolf R; McManus BM  
Corporate Source: UNIV BRITISH COLUMBIA, ST PAULS HOSP, DEPT PATHOL & LAB  
MED, CARDIOVASC RES LAB, MCDONALD RES WING/VANCOUVER/BC V6Z 1Y6/CANADA/  
(REPRINT); UNIV TUBINGEN, INST PATHOL/D-72074 TUBINGEN//GERMANY/  
Journal: VIROLOGY, 1997, V228, N1 (FEB 3), P63-73  
ISSN: 0042-6822 Publication date: 19970203  
Publisher: ACADEMIC PRESS INC JNL-COMP SUBSCRIPTIONS, 525 B ST, STE 1900,  
SAN DIEGO, CA 92101-4495  
Language: English Document Type: ARTICLE (ABSTRACT AVAILABLE)

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DIALOG(R) File 34:SciSearch(R) Cited Ref Sci  
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05540868 Genuine Article#: WF234 Number of References: 45

Title: In vitro mutational and inhibitory analysis of the cis-acting  
translational elements within the 5' untranslated region of  
coxsackievirus B3: Potential targets for antiviral action of  
**antisense** oligomers

Author(s): Yang DC (REPRINT) ; Wilson JE; Anderson DR; Bohunek L; Cordeiro  
C; Kandolf R; McManus BM

Corporate Source: UNIV BRITISH COLUMBIA, ST PAULS HOSP, DEPT PATHOL & LAB  
MED, CARDIOVASC RES LAB, MCDONALD RES WING/VANCOUVER/BC V6Z 1Y6/CANADA/  
(REPRINT); UNIV TUBINGEN, INST PATHOL/D-72074 TUBINGEN//GERMANY/

Journal: VIROLOGY, 1997, V228, N1 (FEB 3), P63-73

ISSN: 0042-6822 Publication date: 19970203

Publisher: ACADEMIC PRESS INC JNL-COMP SUBSCRIPTIONS, 525 B ST, STE 1900,  
SAN DIEGO, CA 92101-4495

Language: English Document Type: ARTICLE  
Geographic Location: CANADA; GERMANY

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Display 4/9/14 (Item 4 from file: 34)  
DIALOG(R) File 34:SciSearch(R) Cited Ref Sci  
(c) 2001 Inst for Sci Info. All rts. reserv.  
Subfile: CC LIFE--Current Contents, Life Sciences  
Journal Subject Category: VIROLOGY

Abstract: The 5' untranslated region (5'UTR) of coxsackievirus B3 (CVB3)  
RNA forms a highly ordered secondary structure that has been implicated  
in controlling initiation of viral translation by internal ribosomal  
entry. To test this hypothesis, synthetic bicistronic RNAs, with all or  
part of the 5'UTR in the intercistronic space, were translated in  
rabbit reticulocyte lysates. In the presence of an upstream cistron,  
the chloramphenicol acetyltransferase gene, designed to block ribosomal  
scanning, the CVB3 5'UTR was capable of directing the internal  
initiation of translation of the downstream reporter gene (Pl),  
confirming the presence of an internal ribosomal entry site (IRES).  
This finding was further supported by the data on predicted secondary  
structures within the 5'UTR. Of special note, analysis of various  
deletion mutants demonstrated that the IRES of CVB3 is located roughly  
at stem-loops G, H, and I spanning nucleotides (nt) 529 and 630. The  
region from nt 1 to 63 (stem-loop A) also appears important, and it may

-more-

?

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DIALOG(R)File 34:SciSearch(R) Cited Ref Sci

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be an essential binding site for translation initiation factors. Based on these findings, in vitro translation inhibition assays using RNA fragments of the 5'UTR as inhibitor were performed. Both **antisense** and sense RNA segments transcribed from these two cis acting regions and the surrounding sequence of the initiation codon AUG showed strong inhibition of viral protein synthesis.

**Antisense** molecules may inhibit translation by blocking ribosome and initiation factor binding within the 5'UTR via specific hybridization to their viral RNA target sequences, while sense sequences may function by competing with viral RNA for ribosomes and/or translation initiation factors. These cis-acting translational elements may serve as potential targets for the antiviral action of oligomers.

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Identifiers--KeyWord Plus(R): ENCEPHALOMYOCARDITIS VIRUS-RNA; INTERNAL RIBOSOME ENTRY; MOUTH-DISEASE VIRUS; NONTRANSLATED REGION; POLIOVIRUS RNA; NONCODING REGION; SECONDARY STRUCTURE; FUNCTIONAL-ANALYSIS; PROTEIN-SYNTHESIS; INITIATION SITE

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? s s1 and fail? and antisense

60916 S1

2791313 FAIL?

141760 ANTISENSE

S5 12 S1 AND FAIL? AND ANTISENSE

? d s5/3/1-12

Display 5/3/1 (Item 1 from file: 5)

DIALOG(R)File 5:BIOSIS Previews(R)

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09321307 BIOSIS NO.: 199497329677

In situ hybridization studies of **hepatitis A** viral RNA in patients with acute **hepatitis A**.

AUTHOR: Taylor Michael(a); Goldin Robert D; Ladva Suresh; Scheuer Peter J; Thomas Howard C

AUTHOR ADDRESS: (a)Dep. Histopathol., St. Mary's Hosp. Med. Sch., Norfolk Place, Paddington, London W2 1PG\*\*UK

JOURNAL: Journal of Hepatology 20 (3):p380-387 1994

ISSN: 0168-8278

DOCUMENT TYPE: Article

RECORD TYPE: Abstract

LANGUAGE: English

- end of record -

?

Display 5/3/2 (Item 1 from file: 55)

DIALOG(R)File 55:BIOSIS Previews(R)

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09321307 BIOSIS NO.: 199497329677

In situ hybridization studies of **hepatitis A** viral RNA in patients with acute **hepatitis A**.

AUTHOR: Taylor Michael(a); Goldin Robert D; Ladva Suresh; Scheuer Peter J; Thomas Howard C

AUTHOR ADDRESS: (a)Dep. Histopathol., St. Mary's Hosp. Med. Sch., Norfolk Place, Paddington, London W2 1PG\*\*UK

JOURNAL: Journal of Hepatology 20 (3):p380-387 1994

ISSN: 0168-8278

DOCUMENT TYPE: Article  
RECORD TYPE: Abstract  
LANGUAGE: English

- end of record -

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Display 5/3/3 (Item 1 from file: 154)  
DIALOG(R) File 154:Medline(R)

08181586 94284584 PMID: 8014450

In situ hybridization studies of **hepatitis A** viral RNA in patients with acute **hepatitis A**.

Taylor M; Goldin RD; Ladva S; Scheuer PJ; Thomas HC  
Department of Histopathology, St. Mary's Hospital Medical School,  
Imperial College of Science, Technology and Medicine, London, UK.

Journal of hepatology (DENMARK) Mar 1994, 20 (3) p380-7, ISSN  
0168-8278 Journal Code: IBS

Languages: ENGLISH

Document type: Journal Article

Record type: Completed

- end of record -

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Display 5/3/4 (Item 1 from file: 155)  
DIALOG(R) File 155:MEDLINE(R)

08181586 94284584 PMID: 8014450

In situ hybridization studies of **hepatitis A** viral RNA in patients with acute **hepatitis A**.

Taylor M; Goldin RD; Ladva S; Scheuer PJ; Thomas HC  
Department of Histopathology, St. Mary's Hospital Medical School,  
Imperial College of Science, Technology and Medicine, London, UK.

Journal of hepatology (DENMARK) Mar 1994, 20 (3) p380-7, ISSN  
0168-8278 Journal Code: IBS

Languages: ENGLISH

Document type: Journal Article

Record type: Completed

- end of record -

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Display 5/3/5 (Item 1 from file: 34)  
DIALOG(R) File 34:SciSearch(R) Cited Ref Sci  
(c) 2001 Inst for Sci Info. All rts. reserv.

03780001 Genuine Article#: QE573 No. References: 156

Title: TREATMENT AND PREVENTION OF CHRONIC VIRAL-HEPATITIS

Author(s): DUSHEIKO GM

Corporate Source: ROYAL FREE HOSP, POND ST/LONDON NW2 2Q3//ENGLAND/; UNIV  
LONDON SCH MED/LONDON NW2 2Q3//ENGLAND/

Journal: PHARMACOLOGY & THERAPEUTICS, 1995, V65, N1, P47-73

ISSN: 0163-7258

Language: ENGLISH Document Type: REVIEW (Abstract Available)

- end of record -

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Display 5/3/6 (Item 2 from file: 34)  
DIALOG(R) File 34:SciSearch(R) Cited Ref Sci  
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03231349    Genuine Article#: NN430    No. References: 19  
Title: IN-SITU HYBRIDIZATION STUDIES OF **HEPATITIS-A** VIRAL-RNA  
      IN PATIENTS WITH ACUTE **HEPATITIS-A**  
Author(s): TAYLOR M; GOLDIN RD; LADVA S; SCHEUER PJ; THOMAS HC  
Corporate Source: UNIV LONDON IMPERIAL COLL SCI TECHNOL & MED, ST MARYS  
      HOSP, SCH MED, DEPT HISTOPATHOL, NORFLOK PL/LONDON W2 1PG//ENGLAND//; UNIV  
      LONDON ROYAL FREE HOSP, DEPT HISTOPATHOL/LONDON//ENGLAND//; UNIV LONDON  
      IMPERIAL COLL SCI TECHNOL & MED, ST MARYS HOSP, SCH MED, DEPT MED/LONDON  
      W2 1PG//ENGLAND/  
Journal: JOURNAL OF HEPATOLOGY, 1994, V20, N3 (MAR), P380-387  
ISSN: 0168-8278  
Language: ENGLISH    Document Type: ARTICLE    (Abstract Available)

- end of record -

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      Display 5/3/7        (Item 1 from file: 73)  
DIALOG(R) File 73: EMBASE  
(c) 2001 Elsevier Science B.V. All rts. reserv.

05676319        EMBASE No: 1994075843  
      In situ hybridization studies of **hepatitis A** viral RNA in  
      patients with acute **hepatitis A**  
      Taylor M.; Goldin R.D.; Ladva S.; Scheuer P.J.; Thomas H.C.  
      Department of Histopathology, St. Mary's Hospital Medical School, Norfolk  
      Place, Paddington, London W2 1PG United Kingdom  
      Journal of Hepatology ( J. HEPATOL. ) (Denmark) 1994, 20/3 (380-387)  
      CODEN: JOHEE    ISSN: 0168-8278  
      DOCUMENT TYPE: Journal; Article  
      LANGUAGE: ENGLISH    SUMMARY LANGUAGE: ENGLISH

- end of record -

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      Display 5/3/8        (Item 1 from file: 94)  
DIALOG(R) File 94: JICST-EPlus  
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02358283    JICST ACCESSION NUMBER: 95A0257289    FILE SEGMENT: JICST-E  
      Forefront of viral hepatitis for clinicians. Molecularly biological  
      approach. Present state and prospect of viral hepatitis.  
      WATANABE AKIHARU (1)  
      (1) Toyama Med. and Pharm. Univ., Fac. of Med.  
      Mod Phys, 1995, VOL.15, NO.1, PAGE.3-7, REF.5  
      JOURNAL NUMBER: X0122ABZ    ISSN NO: 0913-7963  
      UNIVERSAL DECIMAL CLASSIFICATION: 616.3    578.72/.76    575.116  
      LANGUAGE: Japanese        COUNTRY OF PUBLICATION: Japan  
      DOCUMENT TYPE: Journal  
      ARTICLE TYPE: Review article  
      MEDIA TYPE: Printed Publication

- end of record -

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      Display 5/3/9        (Item 1 from file: 98)  
DIALOG(R) File 98: General Sci Abs/Full-Text  
(c) 2001 The HW Wilson Co. All rts. reserv.

04048414    H.W. WILSON    RECORD NUMBER: BGSA99048414    (USE FORMAT 7 FOR  
      FULLTEXT)  
      The unmet challenges of hepatitis C.  
      Di Bisceglie, Adrian M

Bacon, Bruce R  
Scientific American v. 281 no4 (Oct. 1999) p. 80-5  
SPECIAL FEATURES: bibl il ISSN: 0036-8733  
LANGUAGE: English  
COUNTRY OF PUBLICATION: United States  
WORD COUNT: 4233

- end of record -

?

Display 5/3/10 (Item 2 from file: 98)  
DIALOG(R)File 98:General Sci Abs/Full-Text  
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Set	Items	Description
?	s	HAV or hepatitis (n) A
		Processing
		Processing
		Processing
		Processing
	Processed 10 of 37 files ...	
		Processing
		Processing
	Processed 20 of 37 files ...	
		Completed processing all files
	12684	HAV
	610459	HEPATITIS
	61455325	A
	58573	HEPATITIS(N)A
	S1 60910	HAV OR HEPATITIS (N) A
?	s s1 and treat? (5n) fail?	

		Processing
	Processed 10 of 37 files ...	
		Processing
	Processed 20 of 37 files ...	
		Completed processing all files
	60910	S1
	12918210	TREAT?
	2775239	FAIL?
	200572	TREAT?(5N) FAIL?
	S2 195	S1 AND TREAT? (5N) FAIL?
?	s s2 and antisense	
	195	S2
	141760	ANTISENSE
	S3 1	S2 AND ANTISENSE
?	d s3/9/1	

Display 3/9/1 (Item 1 from file: 98)  
 DIALOG(R)File 98:General Sci Abs/Full-Text  
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04048414 H.W. WILSON RECORD NUMBER: BGSA99048414 (THIS IS THE FULLTEXT)  
 The unmet challenges of hepatitis C.  
 Di Bisceglie, Adrian M  
 Bacon, Bruce R  
 Scientific American v. 281 no4 (Oct. 1999) p. 80-5  
 SPECIAL FEATURES: bibl il ISSN: 0036-8733  
 LANGUAGE: English  
 COUNTRY OF PUBLICATION: United States  
 RECORD TYPE: Abstract; Fulltext RECORD STATUS: Corrected or revised  
 record  
 WORD COUNT: 4233

ABSTRACT: Today, almost 4 million people in America suffer from hepatitis C, most of them without knowing it. When researchers first studied viral hepatitis in the 1930s and 1940s, two distinct forms with different patterns of transmission were identified. Later, these viral agents'

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signatures were identified and became known as **hepatitis A** and B, respectively. When new tests became available for the viruses in the 1970s, a new strain was discovered and called non-A, non-B hepatitis, but it was a further 15 years before Michael Houghton and his colleagues at Chiron Corporation identified the hepatitis C virus. The writer discusses how hepatitis C was discovered, how it is transmitted, how it causes chronic liver **failure, treatments** that are currently available for the disease, and the prospects for future treatments.

TEXT:

As recently as the late 1980s few people other than physicians had heard of hepatitis C, a slowly progressing viral infection that over a couple of decades can lead to liver failure or liver cancer. Today the condition is widely recognized as a huge public health concern. Some 1.8 percent of the U.S. adult population, almost four million people, are infected with the hepatitis C virus, most of them without knowing it. The virus is one of the major causes of chronic liver disease, probably

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accounting for even more cases than excessive alcohol use, and is the most common reason for liver transplants. Some 9,000 people die each year in the U.S. from complications of the infection, a number that is expected to triple by 2010. Information about the incidence of hepatitis C in other countries is less reliable, but it is clear that the virus is a major public health problem throughout the world.

Physicians, historians and military leaders have long recognized hepatitis--inflammation of the liver--as a cause of jaundice. This yellow discoloration of the whites of the eyes and skin occurs when the liver fails to excrete a pigment called bilirubin, which then accumulates in the body. In recent decades, however, the diagnosis of hepatitis has progressively improved, and physicians can now distinguish several distinct forms. At least five different viruses can cause the condition, as can drugs and toxins such as alcohol.

Researchers first studied viral hepatitis in the 1930s and 1940s in settings where jaundice was common, such as prisons and mental institutions. They identified two distinct forms with different patterns of

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transmission. One was transmitted by contact with feces of infected individuals and was called infectious hepatitis, or **hepatitis A**. The other appeared to be passed onl